O9634291.O41201

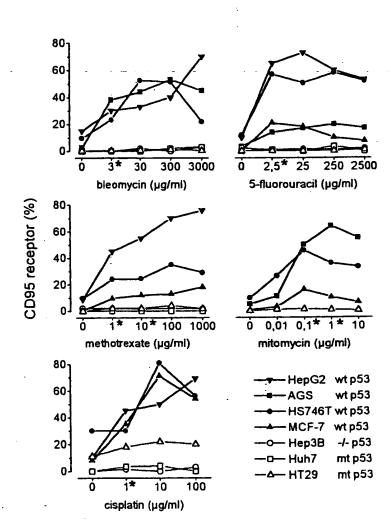


Fig. 1

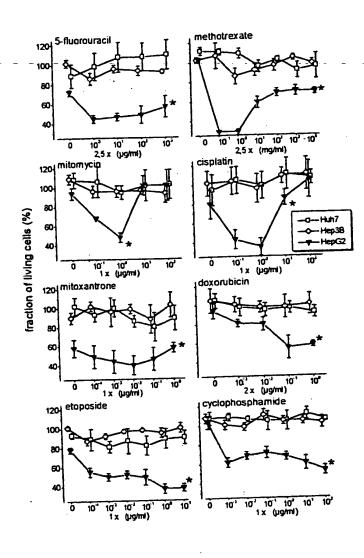


Fig. 2

OGESTES CHIECT

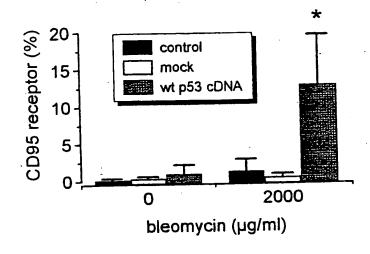


Fig. 3

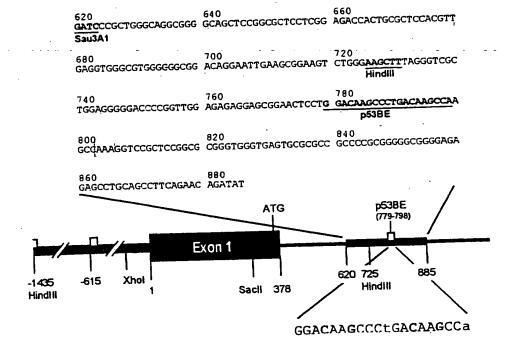


Fig. 4

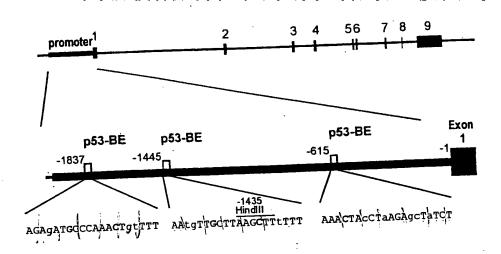


Fig. 5

OSETTOL CHIELL

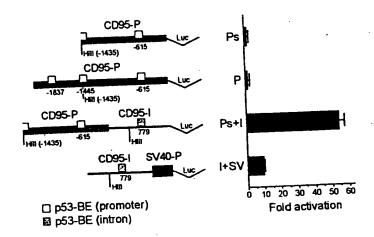


Fig. 6

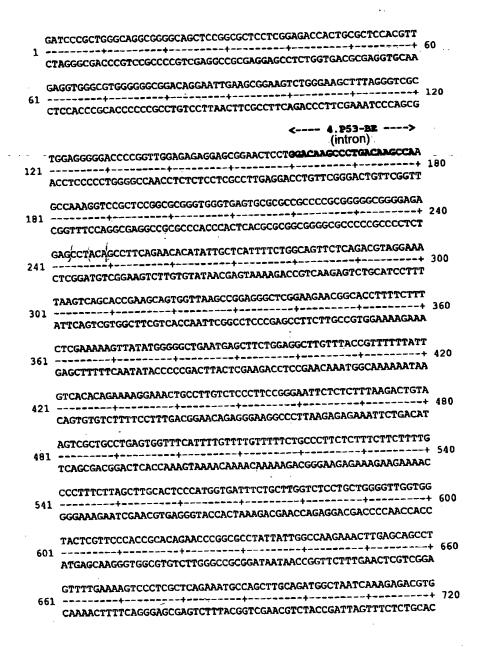


Fig. 7

ogewegt . Chileti

2<sup>nd</sup> half of the 2.p53-BE (promoter)

	480	
	TCGAAAAAACCGATGTAAAAAAATAAACATTTC	
	TAAGTTTAATAATCACTCATCTCACTGGGCTATAATGATAAGTATAAGTAAG	-
31	ATTCAAATTATTAGTGAGTAGAGTGACCCGATATTACTATTCATAATTCATTC	
	CACATATGTGAGTTGCTGGCTTATAATTCACACTCAAGAGATACTGATTTTGTCAATTGT 600	
41	GTGTATACACTCAACGACCGAATATTAAGTGTGAGTTCTCTATGACTAAAACAGTTAACA	
	CCTTTCCCCTTTTTTTCTCTCTCCCTCCTTCCATTCCTTTCCCTTACCTCTCCTTTC	
01	GGAAAGGGGAAAAAAAGAGAAGGGAAGGAAGGTAAGGAAGGAAAGGGAATGGAGAGGGAAAG	
4	CTTCCCTCACACCCCTTTTCCTTCCTTCTTTTTACATTTTTT	
561	GAAGGGACTGTGGGGAAAAGGAAGGAAGAAAATGTAAAAAAAA	
701	ATTTTGGAATAGTTTTAGGATTCAAAAAATTTGCAGAGATAATACAGAGAATGCCCATA	
721	TAAAACCTTATCAAAATCCTAAAGTTTTTTTAAACGTCTCTATTATGTCTCTTACGGGG	
781	TACCATCCTCCTTATCCCACTTCTTTTTGTGTCTATTAGATGCTCAGAGTGTGTGCACAA	
, 01	ATGGTAGGAGGAATAGGGTGAAGAAAAACACAGATAATCTACGAGTCTCACGAG	
841	GGCTGGCACGCCCAGGGTCTTCCTCATGGCACTAACAGTCTACTGAAAGGTGGAACAGAG 900	
	CCGACCGTGCGGGTCCCAGAAGGAGTACCGTGATTGTCAGATGALTTTCCACCTTG	
90	ACAAGCCTATCAACACCTACAAGACTGGTGGTAAGTGCAGTGACAGATGCAAAACACAGG  1	)
	TGTTCGGATAGTTGTGGATGTTCTGACCACCATTCACGTCACTGTCTACGTTTTGTGTCC	
99	GTGATGGAAAGCCCTCAGGAGGGTAACCTAACCTAGATTTGAGGGCCCAAACAGGCTCCA 102	20
	CACTACCTTTCGGGAGTCCTCCCATTGGATTGGATCTAAACTCCCGGGTTTGTCCGAGGT	
102	GAAGAAAATGTCAACTGAGAGGAAGCCTGAAGGATGAACAGTGGGCTAAGCAAAGGGTTA 21	8 (
	21 CTTCTTTTACAGTTGACTCTCCTTCGGACTTCCTACTTGTCACCCGATTCGTTTCCCAAT	

Fig. 8A

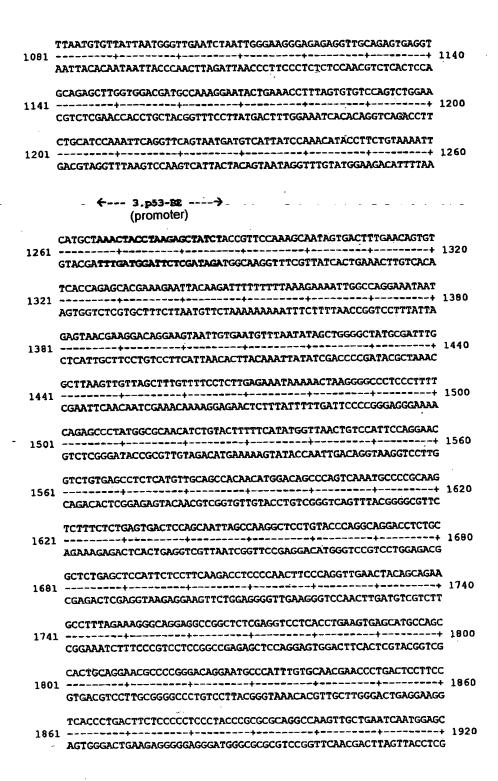
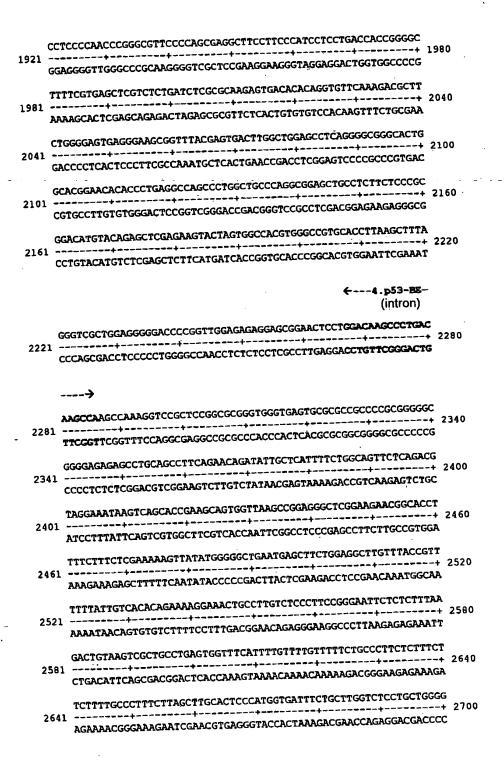


Fig. 8B



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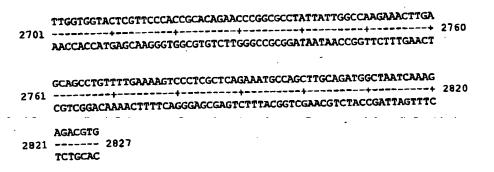


Fig. 8D

<---- 1.p53-BE --TGAGGACTCTCAGGAATATGCTGGTAAAATAAAATAACCTTTACAGATGCCCAAACTGT TTTCCCCAGAACACCAGCATTCATTAGGTGTTCATTCAATAGATTCTTCAAAGGATTCCA 120 AAAGGGGTCTTGTGGTCGTAAGTAATCCACAAGTAAGTTATCTAAGAAGTTTCCTAAGGT AAGGCAAAGAAGTTTGGGGAACAGTATATATAATTACCCAACCCTTTGACATTAGCATAC 121 -----+ 180 TTCCGTTTCTTCAAACCCCTTGTCATATATATTATTGGGTTGGGAAACTGTAATCGTATG TAAGGGCCCTGAGAAGTTTTGGATTAAGAAAGTTTTCAAATTAAAGTAACCCAGAATTTT ATTCCCGGGACTCTTCAAAACCTAATTCTTTCAAAAGTTTAATTTCATTGGGTCTTAAAA CTAAGATTATTTGACCATGAAACATATGTCTCCCCACAAAGCACATATTCCTATCTCCTT 241 -----+ 300 GATTCTAATAAACTGGTACTTTGTATACAGAGGGGTGTTTCGTGTATAAGGATAGAGGAA GAACTTGAGGATAATTAGACGTACGTGGGTAGAGGGTAGGGGAAGGGGGTATGGCATAGA 301 ----- 360 CTTGAACTCCTATTAATCTGCATGCACCCATCTCCCATCCCCTTCCCCCATACCGTATCT <---> 2.p53-E8 ---> 421 -----+ 480 481 -----+ 540 CACATATGTGAGTTGCTGGCTTATAATTCACACTCAAGAGATACTGATTTTGTCAATTGT ----+ 600 GTGTATACACTCAACGACCGAATATTAAGTGTGAGTTCTCTATGACTAAAACAGTTAACA CCTTTCCCCTTTTTTTCTCTCTCCCTCCTTCCATTCCTTCCTTCCCTTACCTCTCCTTTC 

Fig. 9A

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Fig. 9B





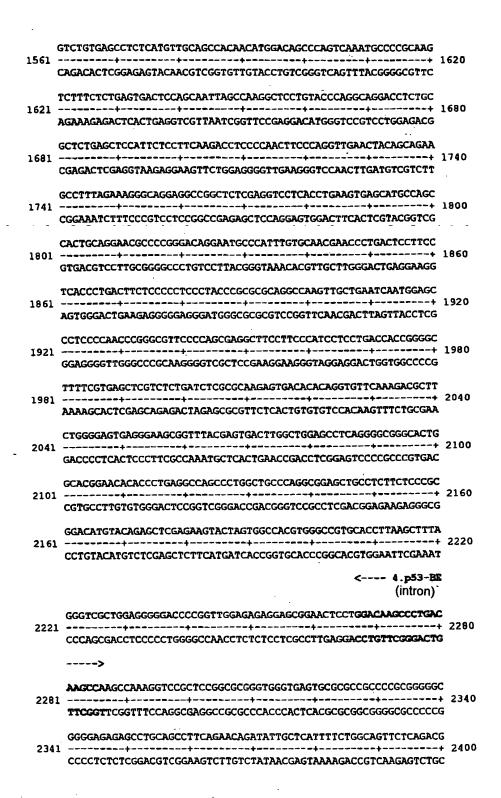


Fig. 9C

2401		2460
	ATCCTTTATTCAGTCGTGGCTTCGTCACCAATTCGGCCTCCCGAGCCTTCTTGCCGTGGA TTTCTTTCTCGAAAAAGTTATATGGGGGCTGAATGAGCTTCTGGAGGCTTGTTTACCGTT	
2461	AAAGAAGAGCTTTTCAATATACCCCCGACTTACTCGAAGACCTCCGAACAAATGGCAA	2520
	TTTTATTGTCACACAGAAAAGGAAACTGCCTTGTCTCCCTTCCGGGAATTCTCTCffTAA	2580
2521	AAAATAACAGTGTGTCTTTTCCTTTGACGGAACAGAGGGAAGGCCCTTAAGAGAGAAATT	
2581	GACTGTAAGTCGCTGCCTGAGTGGTTTCATTTTGTTTTTTTT	2640
2001	CTGACATTCAGCGACGGACTCACCAAAGTAAAACAAAAC	
2641	TCTTTTGCCCTTTCTTAGCTTGCACTCCCATGGTGATTTCTGCTTGGTCTCCTGCTGGGG	2700
	AGAARACGGGAAAGAATCGAACGTGAGGGTACCACTARAGACGAACCAGAGGACGACCCC TTGGTGGTACTCGTTCCCACCGCACAGAACCCGGCGCCTATTATTGGCCAAGAAACTTGA	
2701		2760
	CCACCCTCTTTTGAAAAGTCCCTCGCTCAGAAATGCCAGCTTGCAGATGGCTAATCAAAG	;
2761	CGTCGGACAAAACTTTTCAGGGAGCGAGTCTTTACGGTCGAACGTCTACCGATTAGTTTC	;
2821	AGACGTG 2827 TCTGCAC	

Fig. 9D

<---- 1.p53-88 ---(promoter) TGAGGACTCTCAGGAATATGCTGGTAAAATAAAAATAACCTTTAGAGATGCCCAAACTGT ACTCCTGAGAGTCCTTATACGACCATTTTATTTTTATTGGAAATCTCTACGGGTTTGACA TTTCCCCAGAACACCAGCATTCATTAGGTGTTCATTCAATAGATTCTTCAAAGGATTCCA AAAGGGGTCTTGTGGTCGTAAGTAATCCACAAGTAAGTTATCTAAGAAGTTTCCTAAGGT AAGGCAAAGAAGTTTGGGGAACAGTATATAATTACCCAACCCTTTGACATTAGCATAC \_\_\_\_\_\_ TTCCGTTTCTTCAAACCCCTTGTCATATATATTAATGGGTTGGGAAACTGTAATCGTATG TAAGGGCCCTGAGAAGTTTTGGATTAAGAAAGTTTTCAAATTAAAGTAACCCAGAATTTT ATTCCCGGGACTCTTCAAAACCTAATTCTTTCAAAAGTTTAATTTCATTGGGTCTTAAAA CTAAGATTATTTGACCATGAAACATATGTCTCCCCACAAAGCACATATTCCTATCTCCTT GATTCTAATAAACTGGTACTTTGTATACAGAGGGGTGTTTCGTGTATAAGGATAGAGGAA GAACTTGAGGATAATTAGACGTACGTGGGTAGAGGGTAGGGGAAGGGGGTATGGCATAGA CTTGAACTCCTATTAATCTGCATGCACCCATCTCCCATCCCCTTCCCCCATACCGTATCT <----> (promoter) TGATTGGTAGAAACGG**TTACAACGAATTCGAAAAAA**CCGATGTAAAAAAATAAACATTTC CACATATGTGAGTTGCTGGCTTATAATTCACACTCAAGAGATACTGATTTTGTCAATTGT GTGTATACACTCAACGACCGAATATTAAGTGTGAGTTCTCTATGACTAAAACAGTTAACA CCTTTCCCCTTTTTTTCTCTCTCCCTCCTTCCATTCCTTCTCCCTTACCTCCCTTTC ATTTTGGAATAGTTTTAGGATTTCAAAAAATTTGCAGAGATAATACAGAGAATGCCCCATA TANANCCTTATCANANTCCTANAGTTTTTTANACGTCTCTATTATGTCTCTTACGGGTAT

Fig. 10A

Fig. 10B

TCTGTGAGCCTCTCATGTTGCAGCCACAACATGGACAGCCCAGTCAAATGCCCCGCAAGT

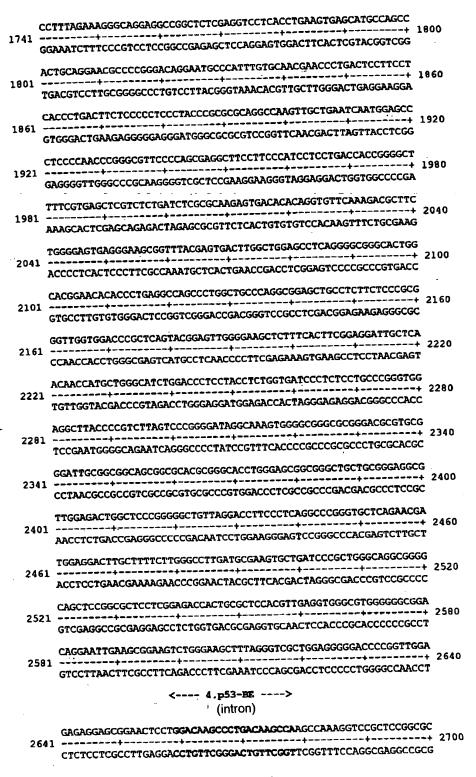


Fig. 10C

Fig. 10D

### Fig. 11A

variations in the p53 binding region of figure 8

### 1. p1140 IMI

p1140

**GGACAAGCCCTGACAAGCCA** 

p1140 IMI

GGANAAGCCCTGACAAGCCA

1

positions of the mutations (boldface and arrow): 2270 (C-A)

### 2. p1140 IMII

p1140

**GGACAAGCCCTGACAAGCCA** 

pl140 IMII

GGAAAAGCCCTGAAAAGCCA

positions of the mutations (boldface and arrow): 2270 (C→A)

2280 (C→A)

#### 3. p1140 IMIII

p1140

GGACAAGCCCTGACAAGCCA

p1140 IMIII

GGAAAATCCCTGAAAATCCA

positions of the mutations (boldface and arrow): 2270 (C->A)

2273 (G→T)

2280 (C→A)

2283 (G→T)

APPROVED	O G. FIG.		
BY	CLISS	SUBCLASS	
DRAFTSMAN	536	24.1	

# Fig. 11B

### 4. p1140 IMIV

p1140

GGACAAGCCCTGACAAGCCA

p1140 IMIV

GCACAAGCCCTCACAAGCCA

1

positions of the mutations (boldface and arrow): 2268 (G $\rightarrow$ T)

. 🕇

2278 (C→A)

DOGS+EG1.O+1EG1

## Fig. 12A

variations in the p53 binding regions of figure 9

#### 1. p1141 IMIII

p1141

**GGACAAGCCCTGACAAGCCA** 

pl141 IMIII

GGAAAATCCCTGAAAATCCA

positions of the mutations (boldface and arrow): 2270

2273

2280

2283

2. p1141 1p53

p1141

AGAGATGCCCAAACTGTTTT

pl141 1p53

AGAGATTCCCAAAATGTTTT

1

positions of the mutations (boldface and arrow): 50

57

3. p1141 2p53

p1141

AATGTTGCTTAAGCTTTTTT

p1141 2p53

AATGTT**T**CTTAAG**A**TT**T**TT

1

positions of the mutations (boldface and arrow): 443

APPROVED O G. FIG. CLASS SUBCLASS BY 24.1 DRAFTSMAN

## Fig. 12B

#### 4. p1141 3p53

p1141

AAACTACCTAAGAGCTATCT

p1141 3p53

ACARTACCTAAGAGCTATCT

1 1

positions of the mutations (boldface and arrow): 1268

 $(A \rightarrow C)$ 

1270

 $(C \rightarrow A)$ 

5. p1141 ΔBgl

<---- 1.p53-BB ---->

p1141

AATAACCTTTAGAGATGCCCAAACTGTTTTCCCCAGAACA

p1141ABgl

6. p1141 ΔSpe

<---- 2.p53-BE ---->

p1141

CATCTTTGCCAATGTTGCTTAAGCTTTTTTTGGCTACATTT

p1141∆Bgl

CATCTTTGCCA-----CTAGTGGCTACATTT

7. p1141 ΔMph

<--->

p1141

AATTCATGCTAAACTACCTAAGAGCTATCTACCGTTCCAA AATTCATGCTATGCA-----TACCGTTCCAA

p1141∆Bgl

Fig. 13A

variations in the p53 binding region of figure 10

#### p1142 TAG

mutation of the positions:

2227 (A→T)

2228 (T→A)

#### p1142 IMIII

p1142

**GGACAAGCCCTGACAAGCCA** 

p1142 IMIII

GGAAAATCCCTGAAAATCCA

positions of the mutations (boldface and arrow): 2662 (C→A)

2665 (G→T)

2672 (C→A)

2675 (G→T)

#### . p1142 ∆Bgl

<---- 1.p53-BE ---->

p1142

AATAACCTTTAGAGATGCCCAAACTGTTTTCCCCCAGAACA

p1142∆Bgl

AATAACCTTTA-----GATCTCCCCAGAACA

#### p1142 ∆Spe

<---->

p1142

CATCTTTGCCAATGTTGCTTAAGCTTTTTTGGCTACATTT

p1142ABgl

CATCTTTGCCA-----CTAGTGGCTACATTT

APPROVED O G. FIG.

BY CLASS SUBCLASS 536 211.1 DRAFTSMAN

Fig. 13B

p1142 ∆Mph 5.

p1142 **p1142∆Bgl** 

DOSS+PJI CHIEDI

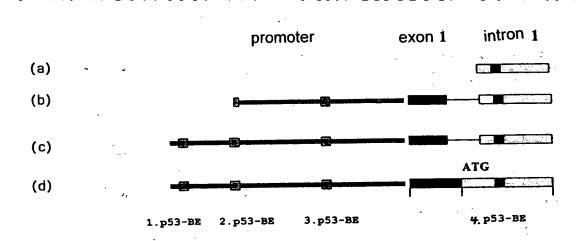


Fig. 14